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SEQUENCE LISTING

<110> Astex Technology Limited  
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Williams, Pamela  
Hamilton, Bruce

<120> Methods of Purification of Cytochrome P450 Proteins

<130> AHBCP6047252

<140> PCT/GB02/02668

<141> 2002-05-30

<160> 84

<170> PatentIn Ver. 2.1

<210> 1

<211> 1428

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: 2C19 (internal deletion, and His tagged) coding sequence.

<400> 1

atggctaaga aaacgagctc taaagggcg cgccctggcc ccactcctc cccagtgatt 60  
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atctatggcc ctgtgttac tctgtatccc ggcctggAAC gcatgggtg gctgcattgg 180  
tatgaagtgg tgaaggaAGC cctgattgtat cttggagagg agtttctgg aagaggccat 240  
ttcccactgg ctgaaagAGC taacagagga tttggaaatcg ttttcagcaa tggaaagaga 300  
tggaaaggaga tccggcgTTT ctccctcatg acgctcgGA attttggat gggaaagagg 360  
agcattgagg accgtgttca agaggaAGC cactgccttgg tggaggagtt gagaAAAacc 420  
aaggcttcac cctgtatcc cacttcatc ctggctgttgc ctccctgcaa tggatctgc 480  
tccattattt tccagaaACG tttcgattat aaagatcAGC aatttcttaa cttgatggaa 540  
aaattgaatg aaaacatcAG gattgtAAcG acccccTGG tccagatATG caataatTTT 600  
cccactatca ttgattattt cccggGAacc cataacaaat tactttttttt cttgctttt 660  
atggaaagtG atatTTTGA gaaagtAAAa gaacaccaAG aatcgatGG aatcaacaAC 720  
cctcgggact ttattgattt ctccctgatc aaaatggAGA aggAAAAGCA aaaccaacAG 780  
tctgaattca ctattgaaaa ctggtaatc actgcagctg acttacttgg agctgggaca 840  
gagacaacaa gcacaacccCT gagatATGCT ctccctctcc tgctgaagca cccagaggTC 900  
acagctaaAG tccagGAAGA gattGAACGT gtcgttggca gaaacccggAG cccctgcATG 960  
caggacaggg gccacatGCC ctacacAGAT gctgtggTC acgaggTCca gagatacATC 1020  
gacccatcc ccaccAGCCT gccccatGCA gtgacccTGT acgttaaattt cagaaactac 1080  
ctcattccca agggcacaac catattaact tccctcactt ctgtgctaca tgacaacaaa 1140  
gaatttccca acccagAGAT gttgaccCT cgtcactttc tgcatGAAGG tggaaatTTT 1200  
aagAAAAGTA actacttcat gccttctca gcaggAAAAC ggatttGTG gggagaggGC 1260  
ctggcccgcA tggagctgtt tttattcctg accttcatt tacagaactt taacctgaaa 1320  
tctctgattt acccaaAGGA ccttgacaca actcctgttG tcaatggatt tgcttctgtc 1380  
ccgccttct accagctctG cttcattccG gtccaccacc accactGA 1428

<210> 2

<211> 475

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Protein  
sequence of 2C19 coded by SEQ ID NO: 1

<400> 2

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15

Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Asp Ile Lys Asp Val Ser  
20 25 30

Lys Ser Leu Thr Asn Leu Ser Lys Ile Tyr Gly Pro Val Phe Thr Leu  
35 40 45

Tyr Phe Gly Leu Glu Arg Met Val Val Leu His Gly Tyr Glu Val Val  
50 55 60

Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly His  
65 70 75 80

Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser  
85 90 95

Asn Gly Lys Arg Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110

Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125

Glu Ala His Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro  
130 135 140

Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160

Ser Ile Ile Phe Gln Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175

Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Arg Ile Val Ser Thr Pro  
180 185 190

Trp Ile Gln Ile Cys Asn Asn Phe Pro Thr Ile Ile Asp Tyr Phe Pro  
195 200 205

Gly Thr His Asn Lys Leu Leu Lys Asn Leu Ala Phe Met Glu Ser Asp  
210 215 220

Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Ile Asn Asn  
225 230 235 240

Pro Arg Asp Phe Ile Asp Cys Phe Leu Ile Lys Met Glu Lys Glu Lys  
245 250 255

Gln Asn Gln Gln Ser Glu Phe Thr Ile Glu Asn Leu Val Ile Thr Ala  
260 265 270

Ala Asp Leu Leu Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285

Tyr Ala Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300

Gln Glu Glu Ile Glu Arg Val Val Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320

Gln Asp Arg Gly His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335

Gln Arg Tyr Ile Asp Leu Ile Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350

Cys Asp Val Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365

Leu Thr Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380

Pro Glu Met Phe Asp Pro Arg His Phe Leu His Glu Gly Gly Asn Phe  
385 390 395 400

Lys Lys Ser Asn Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415

Val Gly Glu Gly Leu Ala Arg Met Glu Leu Phe Leu Phe Leu Thr Phe  
420 425 430

Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Ile Asp Pro Lys Asp Leu  
435 440 445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His  
465 470 475

<210> 3  
<211> 1428  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: 2C19 wild type  
1B

<400> 3  
atggctaaga aaacgagctc taaagggcgg cccgcctggcc ctactcctct cccagtgatt 60  
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atctatggcc ctgtgttac tctgtatccc ggcctggaac gcatgggtggt gctgcatt 180  
tatgaagtgg tgaaggaagc cctgattgtat cttggagagg agtttctgg aagaggccat 240

ttcccaactgg ctgaaagagc taacagagga tttggaatcg ttttcagcaa tggaaagaga 300  
tggaggaga tccggcggtt ctccctcatg acgctgcgga attttggat ggggaagagg 360  
agcattgagg accgtgttca agaggaagcc cgctgcctt tggaggagtt gagaaaaacc 420  
aaagcttcac cctgtgatcc cacttcatc ctgggctgtc ctccctgcaa tgtatctgc 480  
tccatttattt tccagaaacg tttcgattat aaagatcagc aatttcttaa cttgatggaa 540  
aaattgaatg aaaacatcag gattgttaagc acccccttggaa tccagatatg caataattt 600  
cccactatca ttgatttattt cccgggaacc cataacaaat tactaaaaaa cttgctttt 660  
atggaaagtg atattttggaa gaaagtaaaa gaacaccaag aatcgatggaa catcaacaac 720  
cctcgggact ttattgattt cttcctgatc aaaatggaga agggaaagca aaaccaacag 780  
tctgaattca ctattgaaaaa ctggtaatc actgcagctg acttacttgg agctgggaca 840  
gagacaacaa gcacaaccct gagatatgct ctccttctcc tgctgaagca cccagaggc 900  
acagctaaag tccaggaaga gattgaacgt gtcgttggca gaaaccggag cccctgcatg 960  
caggacaggg gccacatgcc ctacacagat gctgtggtgc acgaggcaca gagatacatc 1020  
gacctcatcc ccaccagcct gccccatgca gtgaccctgtc acgttaaattt cagaaactac 1080  
ctcattccca agggcacaac catattaact tccctcactt ctgtgctaca tgacaacaaa 1140  
gaatttccca acccagagat gttgaccct cgtcacttcc tggatgaagg tggaaatttt 1200  
aagaaaagta actacttcat gccttctca gcaggaaaac ggatttgggtt gggagaggc 1260  
ctggcccgca tggagctgtt tttatccctg accttcattt tacagaactt taacctgaaa 1320  
tctctgattt acccaaagga ctttgacaca actcctgtt tcaatggatt tgcttctgtc 1380  
ccgccttctt accagctctg cttcattccct gtccaccacc accactga 1428

<210> 4  
<211> 475  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Translation of  
SEQ ID NO:3

<400> 4  
Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15  
  
Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Asp Ile Lys Asp Val Ser  
20 25 30  
  
Lys Ser Leu Thr Asn Leu Ser Lys Ile Tyr Gly Pro Val Phe Thr Leu  
35 40 45  
  
Tyr Phe Gly Leu Glu Arg Met Val Val Leu His Gly Tyr Glu Val Val  
50 55 60  
  
Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly His  
65 70 75 80  
  
Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser  
85 90 95  
  
Asn Gly Lys Arg Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110  
  
Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125  
  
Glu Ala Arg Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro

130 135 140  
Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160  
Ser Ile Ile Phe Gln Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175  
Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Arg Ile Val Ser Thr Pro  
180 185 190  
Trp Ile Gln Ile Cys Asn Asn Phe Pro Thr Ile Ile Asp Tyr Phe Pro  
195 200 205  
Gly Thr His Asn Lys Leu Leu Lys Asn Leu Ala Phe Met Glu Ser Asp  
210 215 220  
Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Ile Asn Asn  
225 230 235 240  
Pro Arg Asp Phe Ile Asp Cys Phe Leu Ile Lys Met Glu Lys Glu Lys  
245 250 255  
Gln Asn Gln Gln Ser Glu Phe Thr Ile Glu Asn Leu Val Ile Thr Ala  
260 265 270  
Ala Asp Leu Leu Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285  
Tyr Ala Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300  
Gln Glu Glu Ile Glu Arg Val Val Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320  
Gln Asp Arg Gly His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335  
Gln Arg Tyr Ile Asp Leu Ile Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350  
Cys Asp Val Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365  
Leu Thr Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380  
Pro Glu Met Phe Asp Pro Arg His Phe Leu Asp Glu Gly Gly Asn Phe  
385 390 395 400  
Lys Lys Ser Asn Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415  
Val Gly Glu Gly Leu Ala Arg Met Glu Leu Phe Leu Phe Leu Thr Phe  
420 425 430  
Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Ile Asp Pro Lys Asp Leu

435

440

445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His His  
465 470 475

<210> 5  
<211> 1443  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: 2D6 encoding  
nucleic acid

<400> 5

atggctaaaa aaacctcttc taaaggccga ccggccggtc cgctgccgt gccaggcctg 60  
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cgttcgggt acgtgttctc tctcagactg gcttggaccc cggttgttgc tctgaacgg 180  
ctggctgctg ttgcgaaacg tctggttacc cacggtaag acaccgctga ccgtccgccc 240  
gtcccgatca cccagatcct gggttttggg ccgcgttccc aagggtttt cctggctcgt 300  
tacggaccgg cttggcgtga acagcgtcgt ttctctgttt ctaccctgcg taacctggg 360  
ctgggtaaaa aatctctgga acagtgggtt accgaagaag ctgcatgcct gtgcgtcgt 420  
ttcgcttaacc actctggcgt tccgttccgt ccgaacggc tgctggacaa agctgtttt 480  
aacgttatcg cttctctgac ctgcggccgc cggttcaat acgacgaccc gcgttccctg 540  
cgctctgtgg acctggctca ggaagggtctg aaagaggagt ctggttccct gcgtgaagtt 600  
ctgaacgctg ttccgggttct gctgcacatc ccagctctgg ctggtaaagt tctgcgttcc 660  
cagaaaagcat tcctgacccca gctggacgaa ctgctgaccg aacaccgtat gacctgggac 720  
ccggctcagc cgccacgtga cctgaccgaa gctttccctgg ctgaaatgga aaaagctaaa 780  
ggtaaccggg aatctctttt caacgatgaa aatctgcgt a tgcgttgc tgacctgttc 840  
tccgcgggta tggttaccac ctctaccacc ctggcttggg gtctgctgat gatgatcctg 900  
caccggatg tacagcgtcg tggcgtacg gaaatcgacg acgttattgg ccagggtcgt 960  
cgccggaaa tgggtgacca ggctcacatcg ccgtacacca ccgtctttat ccacgaagtt 1020  
cagcgcttcg gtgacatcg tccgtgggtt atgaccacca tgacctctcg tgacatcgaa 1080  
gttcagggtt tccgtatccc gaaagggtacc accctgatca ccaacctgtc ttctgttctg 1140  
aaagacgaag ctgtttgggaaa aaaaaccgttc cggttccatc cggaaacactt cctggacgct 1200  
cagggtaact tcgttaaacc ggaaggcttc ctggccgttct ctgctggcgt tcgtgcttgc 1260  
ctgggtgaaac cgctggctcg tatggaactg ttccgttct tcacctctct gctgcagcac 1320  
ttctctttct ctgttccgac cggtcagccg cgccgtctc accacgggtt tttcgcttcc 1380  
ctggttctc cgtctccgta cgaactgtgc gctgtccgc gtggagctca ccaccaccac 1440  
tga 1443

<210> 6  
<211> 480  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Translation of  
SEQ ID NO: 5

<400> 6

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Leu Pro

1 5 10 15

Leu Pro Gly Leu Gly Asn Leu Leu His Val Asp Phe Gln Asn Thr Pro  
20 25 30

Tyr Cys Phe Asp Gln Leu Arg Arg Arg Phe Gly Asp Val Phe Ser Leu  
35 40 45

Gln Leu Ala Trp Thr Pro Val Val Val Leu Asn Gly Leu Ala Ala Val  
50 55 60

Arg Glu Ala Leu Val Thr His Gly Glu Asp Thr Ala Asp Arg Pro Pro  
65 70 75 80

Val Pro Ile Thr Gln Ile Leu Gly Phe Gly Pro Arg Ser Gln Gly Val  
85 90 95

Phe Leu Ala Arg Tyr Gly Pro Ala Trp Arg Glu Gln Arg Arg Phe Ser  
100 105 110

Val Ser Thr Leu Arg Asn Leu Gly Leu Gly Lys Lys Ser Leu Glu Gln  
115 120 125

Trp Val Thr Glu Glu Ala Ala Cys Leu Cys Ala Ala Phe Ala Asn His  
130 135 140

Ser Gly Arg Pro Phe Arg Pro Asn Gly Leu Leu Asp Lys Ala Val Ser  
145 150 155 160

Asn Val Ile Ala Ser Leu Thr Cys Gly Arg Arg Phe Glu Tyr Asp Asp  
165 170 175

Pro Arg Phe Leu Arg Leu Leu Asp Leu Ala Gln Glu Gly Leu Lys Glu  
180 185 190

Glu Ser Gly Phe Leu Arg Glu Val Leu Asn Ala Val Pro Val Leu Leu  
195 200 205

His Ile Pro Ala Leu Ala Gly Lys Val Leu Arg Phe Gln Lys Ala Phe  
210 215 220

Leu Thr Gln Leu Asp Glu Leu Leu Thr Glu His Arg Met Thr Trp Asp  
225 230 235 240

Pro Ala Gln Pro Pro Arg Asp Leu Thr Glu Ala Phe Leu Ala Glu Met  
245 250 255

Glu Lys Ala Lys Gly Asn Pro Glu Ser Ser Phe Asn Asp Glu Asn Leu  
260 265 270

Arg Ile Val Val Ala Asp Leu Phe Ser Ala Gly Met Val Thr Thr Ser  
275 280 285

Thr Thr Leu Ala Trp Gly Leu Leu Leu Met Ile Leu His Pro Asp Val  
290 295 300

Gln Arg Arg Val Gln Gln Glu Ile Asp Asp Val Ile Gly Gln Val Arg

305	310	315	320
Arg Pro Glu Met Gly Asp Gln Ala His Met Pro Tyr Thr Thr Ala Val			
325	330	335	
Ile His Glu Val Gln Arg Phe Gly Asp Ile Val Pro Leu Gly Met Thr			
340	345	350	
His Met Thr Ser Arg Asp Ile Glu Val Gln Gly Phe Arg Ile Pro Lys			
355	360	365	
Gly Thr Thr Leu Ile Thr Asn Leu Ser Ser Val Leu Lys Asp Glu Ala			
370	375	380	
Val Trp Glu Lys Pro Phe Arg Phe His Pro Glu His Phe Leu Asp Ala			
385	390	395	400
Gln Gly His Phe Val Lys Pro Glu Ala Phe Leu Pro Phe Ser Ala Gly			
405	410	415	
Arg Arg Ala Cys Leu Gly Glu Pro Leu Ala Arg Met Glu Leu Phe Leu			
420	425	430	
Phe Phe Thr Ser Leu Leu Gln His Phe Ser Phe Ser Val Pro Thr Gly			
435	440	445	
Gln Pro Arg Pro Ser His His Gly Val Phe Ala Phe Leu Val Ser Pro			
450	455	460	
Ser Pro Tyr Glu Leu Cys Ala Val Pro Arg Gly Ala His His His His			
465	470	475	480

<210> 7  
 <211> 1458  
 <212> DNA  
 <213> Artificial Sequence

<220>  
 <223> Description of Artificial Sequence: 3A4

<400> 7  
 atggcatacg gtactcattc acatggctcg tttaaaaaac tgggaattcc agggccac 60  
 cctctgcctt ttttggaaaa tattttgtcc taccataagg gctttgtat gtttgacatg 120  
 gaatgtcata aaaagtatgg aaaagtgtgg ggctttatg atggtaaca gcctgtgctg 180  
 gctatcacag atcctgacat gatcaaaaca gtgctagtga aagaatgtta ttctgtcttc 240  
 acaaaccgga ggcctttgg tccagtggta tttatggaaa gtgccatctc tatagctgag 300  
 gatgaagaat ggaagagatt acgatcattt ctgtctccaa ccttcaccag tggaaaactc 360  
 aaggagatgg tccctatcat tgcccagtat ggagatgtgt tggtgagaaa tctgaggcgg 420  
 gaagcagaga caggcaagcc tgtcacctt aaagacgtct ttggggccta cagcatggat 480  
 gtgatcacta gcacatcatt tggagtgaac atcgactctc tcaacaatcc acaagacccc 540  
 tttgtggaaa acaccaagaa gcttttaaga tttgatttt tggatccatt ctttctctca 600  
 ataacagtct ttccatttcct catcccaatt cttgaagtat taaatatctg tgtgtttcca 660  
 agagaagtta caaattttt aagaaaatct gtaaaaagga tgaaagaaag tcgcctcgaa 720

gatacacaaa agcaccgagt ggatttcctt cagctgatga ttgactctca gaattcaaaa 780  
gaaactgagt cccacaaagc tctgtccgat ctggagctcg tggcccaatc aattatctt 840  
attttgctg gctatgaaac cacgagcagt gttctctcct tcattatgta tgaactggcc 900  
actcaccctg atgtccagca gaaactgcag gaggaaattt atgcagttt acccaataag 960  
gcaccaccca cctatgatac tggctacag atggagttt tgcacatggg ggtgaatgaa 1020  
acgctcagat tattccaaat tgctatgaga cttgagaggg tctgaaaaaa agatgtttag 1080  
atcaatggga tggtcattcc caaagggggtg gtggatgaa ttccaagcta tgctttcac 1140  
cgtgacccaa agtactggac agagcctgag aagttcctcc ctgaaagatt cagcaagaag 1200  
aacaaggaca acatagatcc ttacatatac acacccttg gaagtggacc cagaaactgc 1260  
attggcatga ggtttgctct catgaacatg aaacttgctc taatcagagt ctttcagaac 1320  
ttctcattca aacccctgaa agaaaacacag atcccccgtaa aatataagctt aggaggactt 1380  
cttcaaccag aaaaacccgt tggtctaaag gttgagtcaa gggatggcac cgtaagtgga 1440  
gcccaccatc accattga 1458

<210> 8  
<211> 485  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: 3A4

<400> 8  
Met Ala Tyr Gly Thr His Ser His Gly Leu Phe Lys Lys Leu Gly Ile  
1 5 10 15

Pro Gly Pro Thr Pro Leu Pro Phe Leu Gly Asn Ile Leu Ser Tyr His  
20 25 30

Lys Gly Phe Cys Met Phe Asp Met Glu Cys His Lys Lys Tyr Gly Lys  
35 40 45

Val Trp Gly Phe Tyr Asp Gly Gln Gln Pro Val Leu Ala Ile Thr Asp  
50 55 60

Pro Asp Met Ile Lys Thr Val Leu Val Lys Glu Cys Tyr Ser Val Phe  
65 70 75 80

Thr Asn Arg Arg Pro Phe Gly Pro Val Gly Phe Met Lys Ser Ala Ile  
85 90 95

Ser Ile Ala Glu Asp Glu Glu Trp Lys Arg Leu Arg Ser Leu Leu Ser  
100 105 110

Pro Thr Phe Thr Ser Gly Lys Leu Lys Glu Met Val Pro Ile Ile Ala  
115 120 125

Gln Tyr Gly Asp Val Leu Val Arg Asn Leu Arg Arg Glu Ala Glu Thr  
130 135 140

Gly Lys Pro Val Thr Leu Lys Asp Val Phe Gly Ala Tyr Ser Met Asp  
145 150 155 160

Val Ile Thr Ser Thr Ser Phe Gly Val Asn Ile Asp Ser Leu Asn Asn  
165 170 175

Pro Gln Asp Pro Phe Val Glu Asn Thr Lys Lys Leu Leu Arg Phe Asp  
180 185 190

Phe Leu Asp Pro Phe Phe Leu Ser Ile Thr Val Phe Pro Phe Leu Ile  
195 200 205

Pro Ile Leu Glu Val Leu Asn Ile Cys Val Phe Pro Arg Glu Val Thr  
210 215 220

Asn Phe Leu Arg Lys Ser Val Lys Arg Met Lys Glu Ser Arg Leu Glu  
225 230 235 240

Asp Thr Gln Lys His Arg Val Asp Phe Leu Gln Leu Met Ile Asp Ser  
245 250 255

Gln Asn Ser Lys Glu Thr Glu Ser His Lys Ala Leu Ser Asp Leu Glu  
260 265 270

Leu Val Ala Gln Ser Ile Ile Phe Ile Phe Ala Gly Tyr Glu Thr Thr  
275 280 285

Ser Ser Val Leu Ser Phe Ile Met Tyr Glu Leu Ala Thr His Pro Asp  
290 295 300

Val Gln Gln Lys Leu Gln Glu Glu Ile Asp Ala Val Leu Pro Asn Lys  
305 310 315 320

Ala Pro Pro Thr Tyr Asp Thr Val Leu Gln Met Glu Tyr Leu Asp Met  
325 330 335

Val Val Asn Glu Thr Leu Arg Leu Phe Pro Ile Ala Met Arg Leu Glu  
340 345 350

Arg Val Cys Lys Lys Asp Val Glu Ile Asn Gly Met Phe Ile Pro Lys  
355 360 365

Gly Val Val Val Met Ile Pro Ser Tyr Ala Leu His Arg Asp Pro Lys  
370 375 380

Tyr Trp Thr Glu Pro Glu Lys Phe Leu Pro Glu Arg Phe Ser Lys Lys  
385 390 395 400

Asn Lys Asp Asn Ile Asp Pro Tyr Ile Tyr Thr Pro Phe Gly Ser Gly  
405 410 415

Pro Arg Asn Cys Ile Gly Met Arg Phe Ala Leu Met Asn Met Lys Leu  
420 425 430

Ala Leu Ile Arg Val Leu Gln Asn Phe Ser Phe Lys Pro Cys Lys Glu  
435 440 445

Thr Gln Ile Pro Leu Lys Leu Ser Leu Gly Gly Leu Leu Gln Pro Glu  
450 455 460

Lys Pro Val Val Leu Lys Val Glu Ser Arg Asp Gly Thr Val Ser Gly  
465 470 475 480

Ala His His His His  
485

<210> 9  
<211> 9  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Motif

<400> 9  
Ala Lys Lys Thr Ser Ser Lys Gly Arg  
1 5

<210> 10  
<211> 10  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Leader  
sequence

<400> 10  
Met Ala Lys Lys Thr Ser Ser Lys Gly Arg  
1 5 10

<210> 11  
<211> 13  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: N-terminal  
sequence of 3A4

<400> 11  
Met Ala Tyr Gly Thr His Ser His Gly Leu Phe Lys Lys  
1 5 10

<210> 12  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence:  
Oligonucleotide

<400> 12  
caagaggaag cccgctgcct tgtggaggag

<210> 13  
<211> 30  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence:  
Oligonucleotide

<400> 13  
ctcctccaca aggcagcggg ctccctttg 30

<210> 14  
<211> 37  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence:  
Oligonucleotide

<400> 14  
ccctcgac tttctggatg aaggtggaaa ttttaag 37

<210> 15  
<211> 37  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence:  
Oligonucleotide

<400> 15  
cttaaaatccacccat ccagaaatgt acgaggg 37

<210> 16  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 16  
catatggcta aaaaaacctc ttctaaaggc cgaccgcccc gtccgctgcc 50

<210> 17  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 17  
gctgccaggc ctgggtaacc tgctgcatgt ggacttccag aacaccccgta 50

<210> 18  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 18  
actgcttcga ccagctgcgt cgtcgttcg gtgacgtgtt ctctctgcag 50

<210> 19  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 19  
ctggcttggaa ccccggttgt tttctgaac ggtctggctg ctgttcgcga 50

<210> 20  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 20  
agctctggtt acccacggtg aagacaccgc tgaccgtccg ccggtcccga 50

<210> 21  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 21  
tcacccagat cctgggtttt ggtccgcgtt cccaaagggtgt tttcctggct 50

<210> 22  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 22  
cgttacggac cggcttggcg tgaacagcgt cgtttctctg tttctaccct 50

<210> 23  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 23  
gcgtaacctg ggtctggta aaaaatctct ggaacagtgg gttaccgaag 50

<210> 24  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 24  
aagctgcattt cctgtgcgtt gcttcgttta accactctgg tcgtccgttc 50

<210> 25  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 25  
cggtccgaacg gtctgctgga caaagctgtt tctaacgtta tcgtttctct 50

<210> 26

<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 26  
gacctgcggc cgccgttcg aatacgacga cccgcgttcc tgcgtctgc 50

<210> 27  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 27  
tggacctggc tcaggaaggt ctgaaagagg agtctggttt cctgcgtgaa 50

<210> 28  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 28  
gttctgaacg ctgttccggc tctgctgcac atcccagctc tggctggtaa 50

<210> 29  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 29  
agtctgcgt ttccagaaag cattcctgac ccagctggac gaactgctga 50

<210> 30  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 30  
ccgaacacccg tatgacctgg gacccggctc agccgccacg tgacctgacc 50

<210> 31  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 31  
gaagctttcc tggctgaaat ggaaaaagct aaaggttaacc cggaatcttc 50

<210> 32  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 32  
tttcaacgat gaaaatctgc gtatcggtt tgctgacctg ttctccgccc 50

<210> 33  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 33  
gtatggttac cacctctacc accctggctt ggggtctgct gctgatgatc 50

<210> 34  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 34  
ctgcacccgg atgtacagcg tcgtgttcag cagggaaatcg acgacgttat 50

<210> 35  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 35  
tggccaggtt cgtcgccgg aatgggtga ccaggctcac atgccgtaca 50

<210> 36  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 36  
ccaccgctgt tatccacgaa gttcagcgct tcggtgacat cgttccgctg 50

<210> 37  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 37  
ggtatgaccc acatgaccc tcgtgacatc gaagttcagg gtttccgtat 50

<210> 38  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 38  
cccgaaaggt accaccctga tcaccaacct gtcttctgtt ctgaaagacg 50

<210> 39  
<211> 50  
<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 39

aagctgttg ggaaaaaccg ttccgttcc atccggaaca cttcctggac

50

<210> 40

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 40

gctcagggtc acttcgttaa accggaagct ttcctgccgt tctctgctgg

50

<210> 41

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 41

tcgtcgtgct tgcctgggtg aaccgctggc tcgtatggaa ctgttcctgt

50

<210> 42

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 42

tcttcacctc tctgctgcag cacttctctt tctctgttcc gaccggtcag

50

<210> 43

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 43  
ccgcgtccgt ctcaccacgg ttttcgct ttcctggttt ctccgtctcc 50

<210> 44  
<211> 77  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 44  
gtcgactcag tgggggtt gagctccacg cgaaacacgcg cacagttcgt acggagacgg 60  
agaaaccagg aaagcga 77

<210> 45  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 45  
aacaccgtg gtgagacgga cggcgtgac cggtcggAAC agagaaagag 50

<210> 46  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 46  
aagtgcgtca gcagagaggt gaagaacagg aacagttcca tacgagccag 50

<210> 47  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 47  
cggttacccc aggcaagcac gacgaccagc agagaacggc aggaaagctt 50

<210> 48  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 48  
ccggttaac gaagtgaccc tgagcgtcca ggaagtgttc cggatggaaa 50

<210> 49  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 49  
cggaacggtt tttcccaaac agttcgtct ttcagaacag aagacaggtt 50

<210> 50  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 50  
ggtgatcagg gtggcacctt tcgggatacg gaaaccctga acttcgatgt 50

<210> 51  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 51  
cacgagaggt catgtgggtc ataccagcg gaacgatgtc accgaagcgc 50

<210> 52  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 52  
tgaacttcgtt ggataacagc ggtgggtgtac ggcatgtgag cctggtcacc 50

<210> 53  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 53  
catttccggc cgacgaacct ggccaataac gtcgtcgatt tcctgctgaa 50

<210> 54  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 54  
cacgacgctg tacatccggg tgcaggatca tcagcagcag accccaagcc 50

<210> 55  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 55  
agggtggtag aggtggtaac catacccgcg gagaacaggt cagcaacaac 50

<210> 56  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 56  
gatacgcaga ttttcatcgt tgaaagaaga ttccgggtta ccttttagctt 50

<210> 57  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 57  
tttccatttc agccaggaaa gcttcggtca ggtcacgtgg cggctgagcc 50

<210> 58  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 58  
gggtcccagg tcatacggtg ttcggtcagc agttcgtcca gctgggtcag 50

<210> 59  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 59  
gaatgcttcc tggaaacgca gaactttacc agccagagct gggatgtgca 50

<210> 60  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 60  
gcagaaccgg aacagcgttc agaacttcac gcagggaaacc agactcctct 50

<210> 61

<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 61  
ttcagaccaa cctgagccag gtccagcaga cgcaggaaac gcgggtcg 50  
tc

<210> 62  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 62  
gtattcgaaa cggcgccgc aggtcagaga agcgataacg ttagaaacag 50  
gt

<210> 63  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 63  
ctttgtccag cagaccgttc ggacggaacg gacgaccaga gtggtagcg 50  
gt

<210> 64  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 64  
aaagcagcgc acaggcatgc agttttcg gtaaccact gttccagaga 50  
aa

<210> 65  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 65  
ttttttaccc agacccaggt tacgcaggg agaaacagag aaacgacgct 50

<210> 66  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 66  
gttcacgcca agccggtccg taacgagcca ggaaaacacc ttgggaacgc 50

<210> 67  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 67  
ggacccaaaac ccaggatctg ggtgatcggg accggcggac ggtcagcgg 50

<210> 68  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 68  
gtcttcaccg tggtaacca gagttcgcg aacagcagcc agaccgttca 50

<210> 69  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 69  
gaacaacaac cgggtccaa gccagctgca gagagaacac gtcaccgaaa 50

<210> 70  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 70  
cgacgacgca gctggtcgaa gcagtaacggg gtgttctgga agtccacatg 50

<210> 71  
<211> 50  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Oligo used for  
the 2D6 assembly

<400> 71  
cagcaggtta cccaggcctg gcagcggcag cggaccggc ggtcggcctt 50

<210> 72  
<211> 29  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Primer

<400> 72  
gtaacctggg tctggtaaa aaatctctg 29

<210> 73  
<211> 29  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: Primer

<400> 73  
cagagatttt ttacccagac ccaggttac 29

<210> 74  
<211> 33  
<212> DNA  
<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

<400> 74

ggaattcata tggctctcat cccagacttg gcc

33

<210> 75

<211> 47

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

<400> 75

tgcggtcgac tcaatggta tggtgggctc cacttacggt gccatcc

47

<210> 76

<211> 69

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Primer

<400> 76

ttaacatatg gcatatggta ctcattcaca tggtctgttt aaaaaactgg gaattccagg 60  
gccccacacc 69

<210> 77

<211> 475

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: 2C9-FGloop

<400> 77

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15

Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Gly Ile Lys Asp Ile Ser  
20 25 30

Lys Ser Leu Thr Asn Leu Ser Lys Val Tyr Gly Pro Val Phe Thr Leu  
35 40 45

Tyr Phe Gly Leu Lys Pro Ile Val Val Leu His Gly Tyr Glu Ala Val  
50 55 60

Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly Ile  
65 70 75 80

Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser

85

90

95

Asn Gly Lys Lys Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110

Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125

Glu Ala Arg Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro  
130 135 140

Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160

Ser Ile Ile Phe His Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175

Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Lys Ile Leu Ser Ser Pro  
180 185 190

Trp Ile Gln Val Tyr Asn Asn Phe Pro Ala Leu Leu Asp Tyr Phe Pro  
195 200 205

Gly Thr His Asn Lys Leu Leu Lys Asn Val Ala Phe Met Lys Ser Tyr  
210 215 220

Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Met Asn Asn  
225 230 235 240

Pro Gln Asp Phe Ile Asp Cys Phe Leu Met Lys Met Glu Lys Glu Lys  
245 250 255

His Asn Gln Pro Ser Glu Phe Thr Ile Glu Ser Leu Glu Asn Thr Ala  
260 265 270

Val Asp Leu Phe Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285

Tyr Ala Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300

Gln Glu Glu Ile Glu Arg Val Ile Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320

Gln Asp Arg Ser His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335

Gln Arg Tyr Ile Asp Leu Leu Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350

Cys Asp Ile Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365

Leu Ile Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380

Pro Glu Met Phe Asp Pro His His Phe Leu Asp Glu Gly Gly Asn Phe

385

390

395

400

Lys Lys Ser Lys Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415

Val Gly Glu Ala Leu Ala Gly Met Glu Leu Phe Leu Phe Leu Thr Ser  
420 425 430

Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Val Asp Pro Lys Asn Leu  
435 440 445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His His  
465 470 475

<210> 78

<211> 475

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: 2C9-P220 V60C

<400> 78

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15

Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Gly Ile Lys Asp Ile Ser  
20 25 30

Lys Ser Leu Thr Asn Leu Ser Lys Cys Tyr Gly Pro Val Phe Thr Leu  
35 40 45

Tyr Phe Gly Leu Lys Pro Ile Val Val Leu His Gly Tyr Glu Ala Val  
50 55 60

Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly Ile  
65 70 75 80

Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser  
85 90 95

Asn Gly Lys Lys Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110

Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125

Glu Ala Arg Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro  
130 135 140

Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160

Ser Ile Ile Phe His Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175

Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Lys Ile Leu Ser Ser Pro  
180 185 190

Trp Ile Gln Ile Cys Asn Asn Phe Pro Thr Ile Ile Asp Tyr Phe Pro  
195 200 205

Gly Thr His Asn Lys Leu Leu Lys Asn Val Ala Phe Met Lys Ser Tyr  
210 215 220

Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Met Asn Asn  
225 230 235 240

Pro Gln Asp Phe Ile Asp Cys Phe Leu Met Lys Met Glu Lys Glu Lys  
245 250 255

His Asn Gln Pro Ser Glu Phe Thr Ile Glu Ser Leu Glu Asn Thr Ala  
260 265 270

Val Asp Leu Phe Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285

Tyr Ala Leu Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300

Gln Glu Glu Ile Glu Arg Val Ile Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320

Gln Asp Arg Ser His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335

Gln Arg Tyr Ile Asp Leu Leu Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350

Cys Asp Ile Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365

Leu Ile Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380

Pro Glu Met Phe Asp Pro His His Phe Leu Asp Glu Gly Gly Asn Phe  
385 390 395 400

Lys Lys Ser Lys Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415

Val Gly Glu Ala Leu Ala Gly Met Glu Leu Phe Leu Phe Leu Thr Ser  
420 425 430

Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Val Asp Pro Lys Asn Leu  
435 440 445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His His  
465 470 475

<210> 79  
<211> 475  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: 2C9-P220

<400> 79  
Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15  
  
Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Gly Ile Lys Asp Ile Ser  
20 25 30  
  
Lys Ser Leu Thr Asn Leu Ser Lys Val Tyr Gly Pro Val Phe Thr Leu  
35 40 45  
  
Tyr Phe Gly Leu Lys Pro Ile Val Val Leu His Gly Tyr Glu Ala Val  
50 55 60  
  
Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly Ile  
65 70 75 80  
  
Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser  
85 90 95  
  
Asn Gly Lys Lys Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110  
  
Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125  
  
Glu Ala Arg Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro  
130 135 140  
  
Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160  
  
Ser Ile Ile Phe His Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175  
  
Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Lys Ile Leu Ser Ser Pro  
180 185 190  
  
Trp Ile Gln Ile Cys Asn Asn Phe Pro Thr Ile Ile Asp Tyr Phe Pro  
195 200 205  
  
Gly Thr His Asn Lys Leu Leu Lys Asn Val Ala Phe Met Lys Ser Tyr  
210 215 220  
  
Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Met Asn Asn  
225 230 235 240

Pro Gln Asp Phe Ile Asp Cys Phe Leu Met Lys Met Glu Lys Glu Lys  
245 250 255

His Asn Gln Pro Ser Glu Phe Thr Ile Glu Ser Leu Glu Asn Thr Ala  
260 265 270

Val Asp Leu Phe Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285

Tyr Ala Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300

Gln Glu Glu Ile Glu Arg Val Ile Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320

Gln Asp Arg Ser His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335

Gln Arg Tyr Ile Asp Leu Leu Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350

Cys Asp Ile Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365

Leu Ile Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380

Pro Glu Met Phe Asp Pro His His Phe Leu Asp Glu Gly Gly Asn Phe  
385 390 395 400

Lys Lys Ser Lys Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415

Val Gly Glu Ala Leu Ala Gly Met Glu Leu Phe Leu Phe Leu Thr Ser  
420 425 430

Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Val Asp Pro Lys Asn Leu  
435 440 445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His His  
465 470 475

<210> 80  
<211> 475  
<212> PRT  
<213> Artificial Sequence

<220>  
<223> Description of Artificial Sequence: 2C9-FG  
Loop-K206E

<400> 80

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15

Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Gly Ile Lys Asp Ile Ser  
20 25 30

Lys Ser Leu Thr Asn Leu Ser Lys Val Tyr Gly Pro Val Phe Thr Leu  
35 40 45

Tyr Phe Gly Leu Lys Pro Ile Val Val Leu His Gly Tyr Glu Ala Val  
50 55 60

Lys Glu Ala Leu Ile Asp Leu Gly Glu Glu Phe Ser Gly Arg Gly Ile  
65 70 75 80

Phe Pro Leu Ala Glu Arg Ala Asn Arg Gly Phe Gly Ile Val Phe Ser  
85 90 95

Asn Gly Lys Lys Trp Lys Glu Ile Arg Arg Phe Ser Leu Met Thr Leu  
100 105 110

Arg Asn Phe Gly Met Gly Lys Arg Ser Ile Glu Asp Arg Val Gln Glu  
115 120 125

Glu Ala Arg Cys Leu Val Glu Glu Leu Arg Lys Thr Lys Ala Ser Pro  
130 135 140

Cys Asp Pro Thr Phe Ile Leu Gly Cys Ala Pro Cys Asn Val Ile Cys  
145 150 155 160

Ser Ile Ile Phe His Lys Arg Phe Asp Tyr Lys Asp Gln Gln Phe Leu  
165 170 175

Asn Leu Met Glu Lys Leu Asn Glu Asn Ile Glu Ile Leu Ser Ser Pro  
180 185 190

Trp Ile Gln Val Tyr Asn Asn Phe Pro Ala Leu Leu Asp Tyr Phe Pro  
195 200 205

Gly Thr His Asn Lys Leu Leu Lys Asn Val Ala Phe Met Lys Ser Tyr  
210 215 220

Ile Leu Glu Lys Val Lys Glu His Gln Glu Ser Met Asp Met Asn Asn  
225 230 235 240

Pro Gln Asp Phe Ile Asp Cys Phe Leu Met Lys Met Glu Lys Glu Lys  
245 250 255

His Asn Gln Pro Ser Glu Phe Thr Ile Glu Ser Leu Glu Asn Thr Ala  
260 265 270

Val Asp Leu Phe Gly Ala Gly Thr Glu Thr Thr Ser Thr Thr Leu Arg  
275 280 285

Tyr Ala Leu Leu Leu Leu Lys His Pro Glu Val Thr Ala Lys Val  
290 295 300

Gln Glu Glu Ile Glu Arg Val Ile Gly Arg Asn Arg Ser Pro Cys Met  
305 310 315 320

Gln Asp Arg Ser His Met Pro Tyr Thr Asp Ala Val Val His Glu Val  
325 330 335

Gln Arg Tyr Ile Asp Leu Leu Pro Thr Ser Leu Pro His Ala Val Thr  
340 345 350

Cys Asp Ile Lys Phe Arg Asn Tyr Leu Ile Pro Lys Gly Thr Thr Ile  
355 360 365

Leu Ile Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu Phe Pro Asn  
370 375 380

Pro Glu Met Phe Asp Pro His His Phe Leu Asp Glu Gly Gly Asn Phe  
385 390 395 400

Lys Lys Ser Lys Tyr Phe Met Pro Phe Ser Ala Gly Lys Arg Ile Cys  
405 410 415

Val Gly Glu Ala Leu Ala Gly Met Glu Leu Phe Leu Phe Leu Thr Ser  
420 425 430

Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Val Asp Pro Lys Asn Leu  
435 440 445

Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro Pro Phe Tyr  
450 455 460

Gln Leu Cys Phe Ile Pro Val His His His His  
465 470 475

<210> 81

<211> 494

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: 2C9 wild type  
P450

<400> 81

Met Asp Ser Leu Val Val Leu Val Cys Leu Ser Cys Leu Leu Leu  
1 5 10 15

Leu Ser Leu Trp Arg Gln Ser Ser Gly Arg Gly Lys Leu Pro Pro Gly  
20 25 30

Pro Thr Pro Leu Pro Val Ile Gly Asn Ile Leu Gln Ile Gly Ile Lys  
35 40 45

Asp Ile Ser Lys Ser Leu Thr Asn Leu Ser Lys Val Tyr Gly Pro Val  
50 55 60

Phe Thr Leu Tyr Phe Gly Leu Lys Pro Ile Val Val Leu His Gly Tyr

65	70	75	80
Glu Ala Val Lys Glu Ala Leu Ile Asp Leu Gly		Glu Glu Phe Ser Gly	
85		90	95
Arg Gly Ile Phe Pro Leu Ala Glu Arg Ala Asn Arg	Gly Phe Gly Ile		
100	105		110
Val Phe Ser Asn Gly Lys Lys Trp Lys Glu Ile Arg	Arg Phe Ser Leu		
115	120		125
Met Thr Leu Arg Asn Phe Gly Met Gly Lys Arg Ser	Ile Glu Asp Arg		
130	135		140
Val Gln Glu Glu Ala Arg Cys Leu Val Glu Glu Leu	Arg Lys Thr Lys		
145	150		155
Ala Ser Pro Cys Asp Pro Thr Phe Ile Leu Gly Cys	Ala Pro Cys Asn		
165	170		175
Val Ile Cys Ser Ile Ile Phe His Lys Arg Phe Asp	Tyr Lys Asp Gln		
180	185		190
Gln Phe Leu Asn Leu Met Glu Lys Leu Asn Glu Asn	Ile Lys Ile Leu		
195	200		205
Ser Ser Pro Trp Ile Gln Ile Cys Asn Asn Phe	Ser Pro Ile Ile Asp		
210	215		220
Tyr Phe Pro Gly Thr His Asn Lys Leu Leu Lys Asn	Val Ala Phe Met		
225	230		235
Lys Ser Tyr Ile Leu Glu Lys Val Lys Glu His Gln	Glu Ser Met Asp		
245	250		255
Met Asn Asn Pro Gln Asp Phe Ile Asp Cys Phe	Leu Met Lys Met Glu		
260	265		270
Lys Glu Lys His Asn Gln Pro Ser Glu Phe Thr Ile	Glu Ser Leu Glu		
275	280		285
Asn Thr Ala Val Asp Leu Phe Gly Ala Gly Thr	Glu Thr Thr Ser Thr		
290	295		300
Thr Leu Arg Tyr Ala Leu Leu Leu Leu Lys His	Pro Glu Val Thr		
305	310		320
Ala Lys Val Gln Glu Glu Ile Glu Arg Val Ile Gly	Arg Asn Arg Ser		
325	330		335
Pro Cys Met Gln Asp Arg Ser His Met Pro Tyr	Thr Asp Ala Val Val		
340	345		350
His Glu Val Gln Arg Tyr Ile Asp Leu Leu Pro	Thr Ser Leu Pro His		
355	360		365
Ala Val Thr Cys Asp Ile Lys Phe Arg Asn Tyr	Leu Ile Pro Lys Gly		

370

375

380

Thr Thr Ile Leu Ile Ser Leu Thr Ser Val Leu His Asp Asn Lys Glu  
385 390 395 400

Phe Pro Asn Pro Glu Met Phe Asp Pro His His Phe Leu Asp Glu Gly  
405 410 415

Gly Asn Phe Lys Lys Ser Lys Tyr Phe Met Pro Phe Ser Ala Gly Lys  
420 425 430

Arg Ile Cys Val Gly Glu Ala Leu Ala Gly Met Glu Leu Phe Leu Phe  
435 440 445

Leu Thr Ser Ile Leu Gln Asn Phe Asn Leu Lys Ser Leu Val Asp Pro  
450 455 460

Lys Asn Leu Asp Thr Thr Pro Val Val Asn Gly Phe Ala Ser Val Pro  
465 470 475 480

Pro Phe Tyr Gln Leu Cys Phe Ile Pro Val His His His His  
485 490

<210> 82

<211> 40

<212> PRT

<213> Homo sapiens

<400> 82

Met Asp Ser Leu Val Val Leu Val Leu Cys Leu Ser Cys Leu Leu Leu  
1 5 10 15

Leu Ser Leu Trp Arg Gln Ser Ser Gly Arg Gly Lys Leu Pro Pro Gly  
20 25 30

Pro Thr Pro Leu Pro Val Ile Gly  
35 40

<210> 83

<211> 21

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: 2C9 and 2C19  
truncation

<400> 83

Met Ala Lys Lys Thr Ser Ser Lys Gly Arg Pro Pro Gly Pro Thr Pro  
1 5 10 15

Leu Pro Val Ile Gly  
20

<210> 84  
<211> 40  
<212> PRT  
<213> Homo sapiens

<400> 84  
Met Asp Pro Phe Val Val Leu Val Leu Cys Leu Ser Cys Leu Leu Leu  
1 5 10 15  
Leu Ser Ile Trp Arg Gln Ser Ser Gly Arg Gly Lys Leu Pro Pro Gly  
20 25 30  
Pro Thr Pro Leu Pro Val Ile Gly  
35 40